

Presentation at Biotech Showcase 2025

Melbourne, Australia – Tryptamine Therapeutics Limited ('Tryp' or the 'Company') (ASX: TYP), a clinical-stage biopharmaceutical company, is pleased to provide the following presentation which will be given at renowned investor conference, the Biotech Showcase 2025, in San Francisco on 15 January 2025.

Biotech Showcase is a dedicated investor conference designed to provide private and micro-mid-cap biotechnology companies with the opportunity to present and connect with investors and industry executives. Tryp was selected as one of 400 companies from across the globe to present its recent findings and progress. Presenting at the event will provide the Company with direct access to investor groups that collectively manage over US\$400bn in capital, along with key strategic partners in the life sciences sector.

This announcement has been authorised for release by the Board of Tryptamine Therapeutics Limited.

-ENDS-

About Tryptamine Therapeutics Limited

Tryp Therapeutics is a clinical-stage biopharmaceutical company focused on developing proprietary, novel formulations for the administration of psilocin in combination with psychotherapy to treat diseases with unmet medical needs. Tryp's lead asset, TRP-8803, is a proprietary, scalable and innovative formulation of IV-infused psilocin (the active metabolite of psilocybin) with neuroplastic benefits. It has the potential to alleviate numerous shortcomings of oral psilocybin including: significantly reducing the time to onset of the neuroplastic state, controlling the depth and duration of the neuroplastic experience, and reducing the overall duration of the intervention to a commercially feasible timeframe. The Company has completed a Phase 2a clinical trial for the treatment of binge eating disorder at the University of Florida, which demonstrated an average reduction in binge eating episodes of greater than 80%.

The Company also has also just completed a Phase 2a successful clinical trial for the treatment of fibromyalgia in collaboration with the University of Michigan and has initiated a Phase 2a clinical trial in collaboration with Massachusetts General Hospital for the treatment of abdominal pain and visceral tenderness in patients suffering from irritable bowel syndrome. Each of the studies is utilising TRP-8802 (synthetic, oral psilocybin) to demonstrate clinical benefit in these indications. Where a positive clinical response is demonstrated, subsequent studies are expected to utilise TRP-8803 (IV-infused psilocin), that has the potential to further improve efficacy, safety, and patient experience.

For more information, please visit www.tryptherapeutics.com.

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Risks associated with psilocin

All medicines carry risks and specialist prescribers, such as registered psychiatrists are best placed to assess the suitability of a new medication against a patient's individual circumstances and medical history before proceeding. Adverse effects of psilocybin and similar compounds, such as psilocin, can include temporary increase in blood pressure and a raised heart rate. There may be some risk of psychosis in predisposed individuals. These effects of psilocybin and its derivatives are unlikely at low doses and in the treatment regimens used in psychedelic-assisted psychotherapy and appropriately managed in a controlled environment with direct medical supervision.

Forward-Looking Information

Certain information in this news release, constitutes forward looking information. In some cases, but not necessarily in all cases, forward-looking information can be identified by the use of forward-looking terminology such as "plans", "targets", "expects" or "does not expect", "is expected", "an opportunity exists", "is positioned", "estimates", "intends", "assumes", "anticipates" or "does not anticipate" or "believes", or variations of such words and phrases or state that certain actions, events or results "may", "could", "would", "might", "will" or "will be taken", "occur" or "be achieved". In addition, any statements that refer to expectations, projections or other characterizations of future events or circumstances contain forward-looking information. Statements containing forward-looking information are not historical facts but instead represent management's expectations, estimates and projections regarding future events. Forward-looking information is necessarily based on a number of opinions, assumptions and estimates that, while considered reasonable by Tryp as of the date of this news release, are subject to known and unknown risks, uncertainties, assumptions and other factors that may cause the actual results, level of activity, performance or achievements to be materially different from those expressed or implied by such forward looking information, including but not limited to the factors described in greater detail in the "Risk Factors" section of Tryp's Replacement Prospectus available at www.asx.com.au These factors are not intended to represent a complete list of the factors that could affect Tryp; however, these factors should be considered carefully. There can be no assurance that such estimates and assumptions will prove to be correct. The forward-looking statements contained in this news release are made as of the date of this news release, and Tryp expressly disclaims any obligation to update or alter statements containing any forward-looking information, or the factors or assumptions underlying them, whether as a result of new information, future events or otherwise, except as required by law.



TRYPTAMINE
THERAPEUTICS

Precision in Psychedelic Therapy

Biotech Showcase 2025, San Francisco, USA

ASX : TYP

This presentation has been authorised for release by the Board of Tryptamine Therapeutics Limited

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Psilocybin. Psilocybin is currently a Schedule III drug under the Controlled Drugs and Substances Act, S.C. 1996, c. 19 (the “CDSA”) and it is a criminal offence to possess substances under the CDSA without a prescription. Health Canada has not approved psilocybin as a drug. While the Company is focused on developing products using psilocybin, the Company does not have any direct or indirect involvement with the illegal selling, production or distribution of any substances. The Company does not currently manufacture, store or otherwise handle psilocybin directly and will only do so through agents within laboratory and clinical trial settings conducted within approved regulatory frameworks. The Company’s products that contain psilocybin or other psychedelic compounds will not be commercialized prior to applicable regulatory approval, which will only be granted if clinical evidence of safety and efficacy for the intended uses is successfully developed.

All medicines carry risks and specialist prescribers, such as registered psychiatrists are best placed to assess the suitability of a new medication against a patient’s individual circumstances and medical history before proceeding.

Adverse effects of psilocybin and its derivatives can include temporary increase in blood pressure and a raised heart rate. There may be some risk of psychosis in predisposed individuals. These effects of psilocybin and its derivatives are unlikely at low doses and in the treatment regimens used in psychedelic-assisted psychotherapy and appropriately managed in a controlled environment with direct medical supervision.

Delivering Precision to Neuroplastic Medicine

With a focus on solving:
EATING DISORDERS & CHRONIC PAIN

Tryp is a clinical stage drug development company developing an innovative and scalable IV-infused psilocin solution which can be used with therapy to address unmet medical needs

Infusion refers to the continuous administration of the drug via the intravenous route and provides a clinician with full control over the administration of the solution

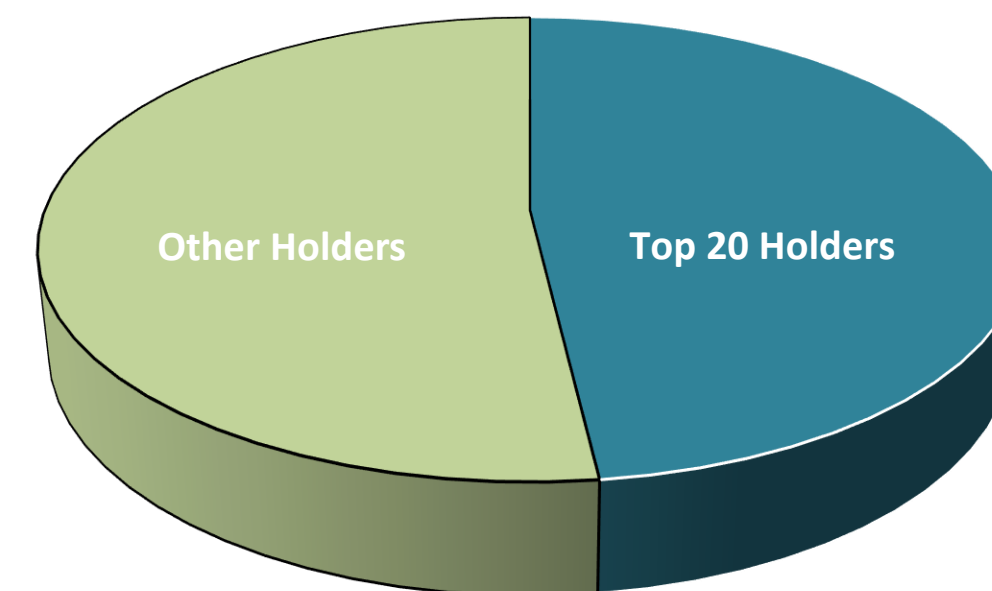


Corporate overview

Snapshot:	
ASX code:	TYP
Shares on issue:	1.227Bn
Market capitalisation: (at \$0.04 per share)	AU\$49.1m
Cash at bank: (as at 30 Sep 2024)	AU\$5.3m*
Debt:	Nil

Board of Directors	
Non-Executive Chairman	Mr. Mark Davies
Chief Executive Officer	Mr. Jason Carroll
Non-Executive Director	Dr. Daniel Tillett
Non-Executive Director	Mr. Chris Ntoumenopoulos
Non-Executive Director	Mr. Gage Jull

*Cash balance at 30 Sept 2024 excludes expected R&D Tax Rebate Incentive [AU\$1.05M] and net capital from Tranche 2 raises [AU\$3.05M]



Major shareholders (at 18 December 2024)	
Dr. William James Garner (<i>co-founder</i>)	15.5%
Citicorp Nominees P/L	7.5%
Mr. Jason Carroll (<i>CEO</i>)	2.9%
BNP Paribas Nominees P/L	2.7%
Mr. Herwig Janssen (<i>HNW/J&J Executive</i>)	2.0%
Top 5:	30.6%
Top 10:	38.7%
Top 20:	47.7%
Top 100:	80.5%

Investment highlights

Focused on the development of a best in class neuroplastegen and precise dosing in patients with neuropsychiatric disorders

A transformative and commercially scalable IP portfolio via the use of IV-infused psilocin (TRP-8803):

- TRP-8803 addresses the limitations of oral dosing – Include improvements in treatment time and the ability to reverse treatment quickly
- Platform has broad applicability and out-licensing potential across multiple indications

Strong clinical trial pipeline (completed and ongoing):

- Three pathfinder Phase 2 clinical trials - Binge Eating Disorder (complete), Fibromyalgia (complete) and Irritable Bowel Syndrome (ongoing)
- Each Phase 2 trial delivered meaningful results into specific indications using oral dosing (TRP-8802)
- Phase 1 healthy human volunteer study using TRP-8803 completed in 14 patients to define optimal infusion rates - TRP-8803 deemed safe by SRC
- Preparation for first active patient clinical study using TRP-8803 in Australia well advanced

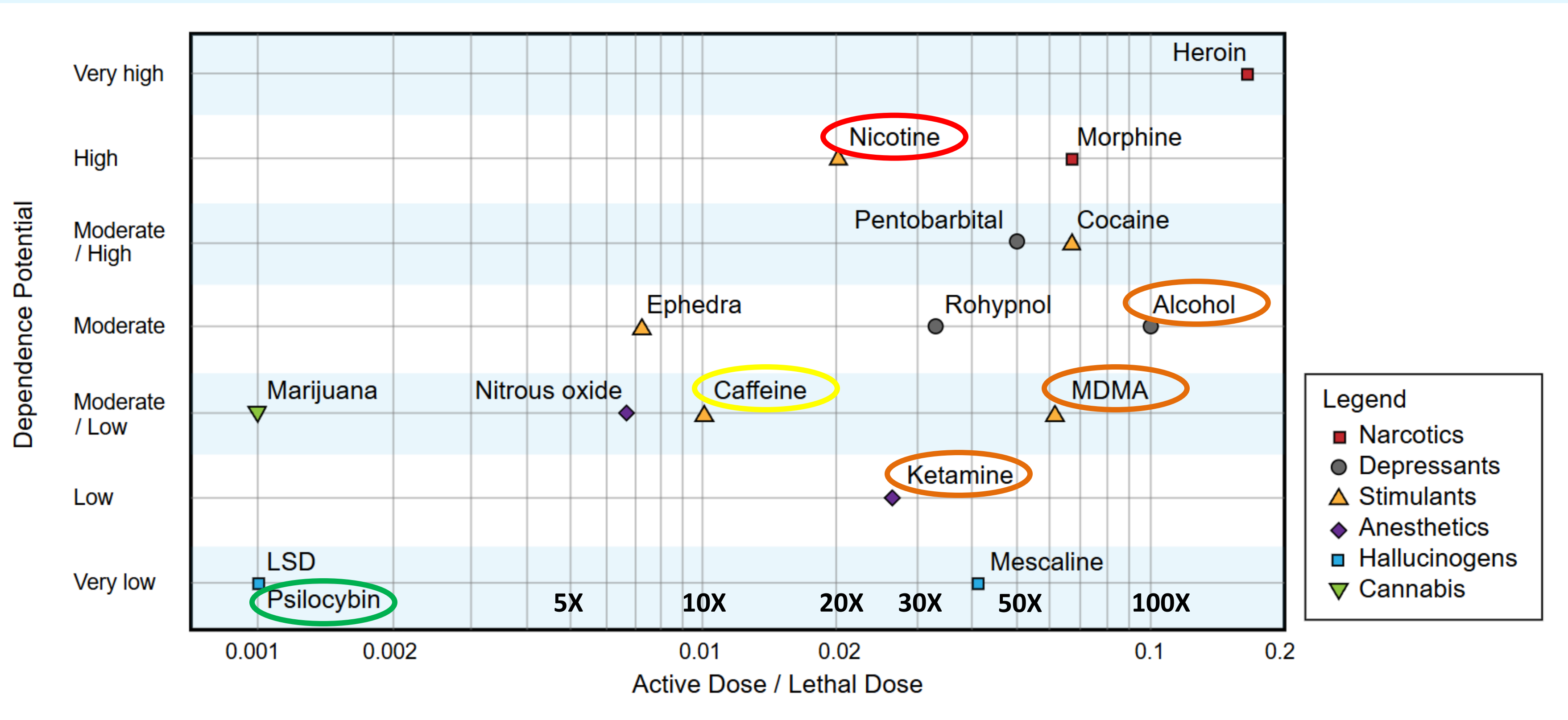
A world-class team, partners and advisory board:

- Board and management include executives with over 20 years' experience in pharmaceutical and drug development
- Scientific Advisory Board lead by sector expert, Robin Carhart-Harris and underpinned by leading Australian professors
- Exceptional research partners including Massachusetts General Hospital, the University of Florida and others

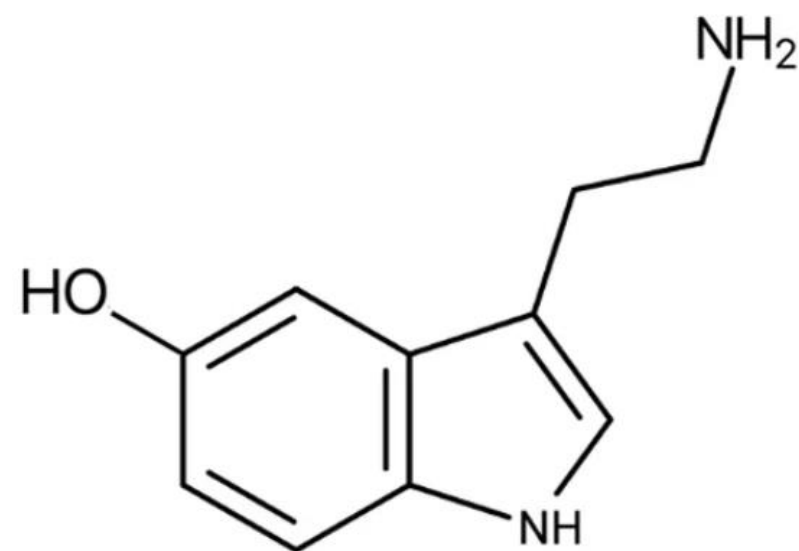
Funded and well placed to capitalise on regulation:

- Australian listed entity allows company to benefit from positive regulatory stance and leverage government R&D incentives
- Recent A\$6m placement with renowned biotechnology investors provides strong financial flexibility

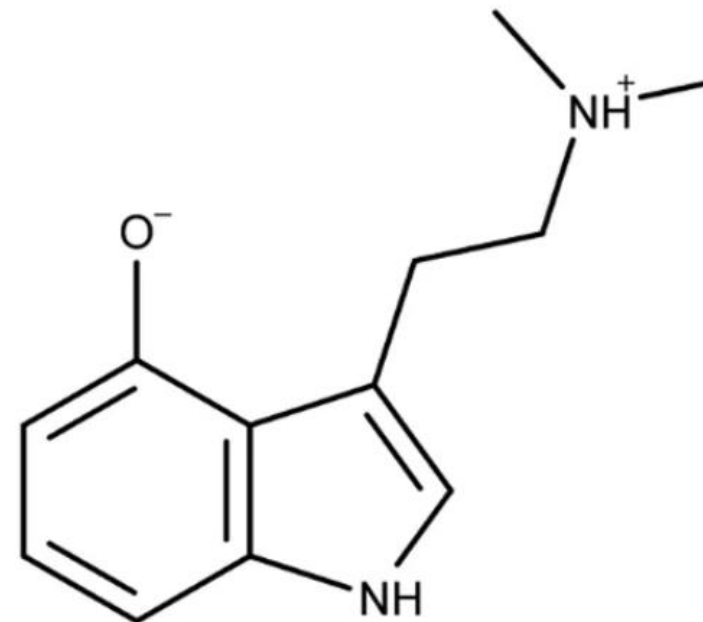
Dispelling myths regarding pharmaceutical safety of Psilocybin



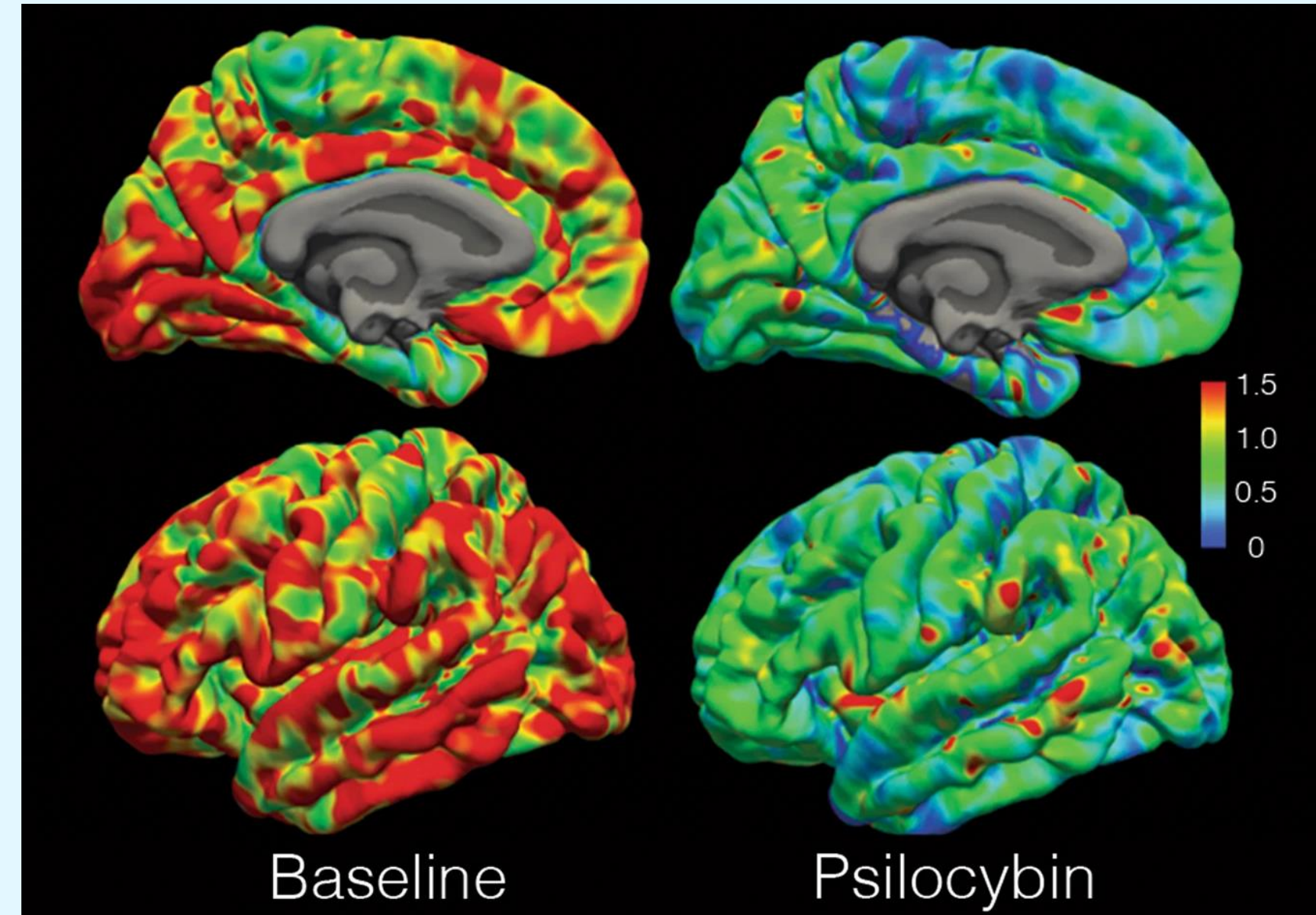
Efficacy in targeting serotonin 5-HT_{2A} receptors



Serotonin
(5-Hydroxytryptamine)



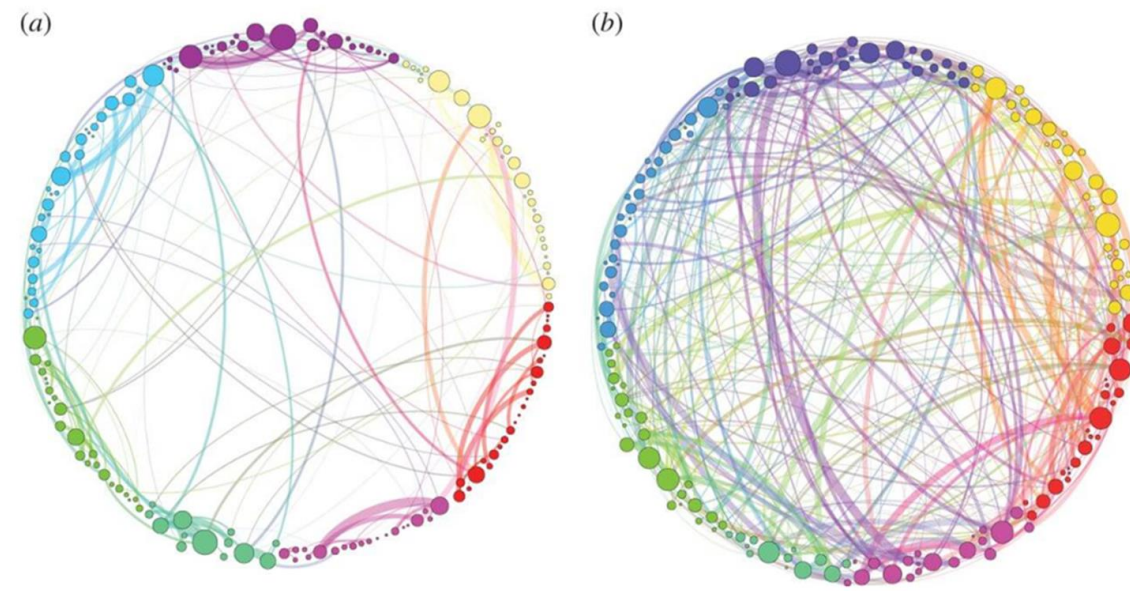
Psilocin
(4-Hydroxy-N,N-Dimethyltryptamine)



Psilocin molecules activate the serotonin 5HT_{2A} receptor due to structural similarity between psilocin & serotonin.

Psilocin occupancy of 5-HT_{2A} Receptors

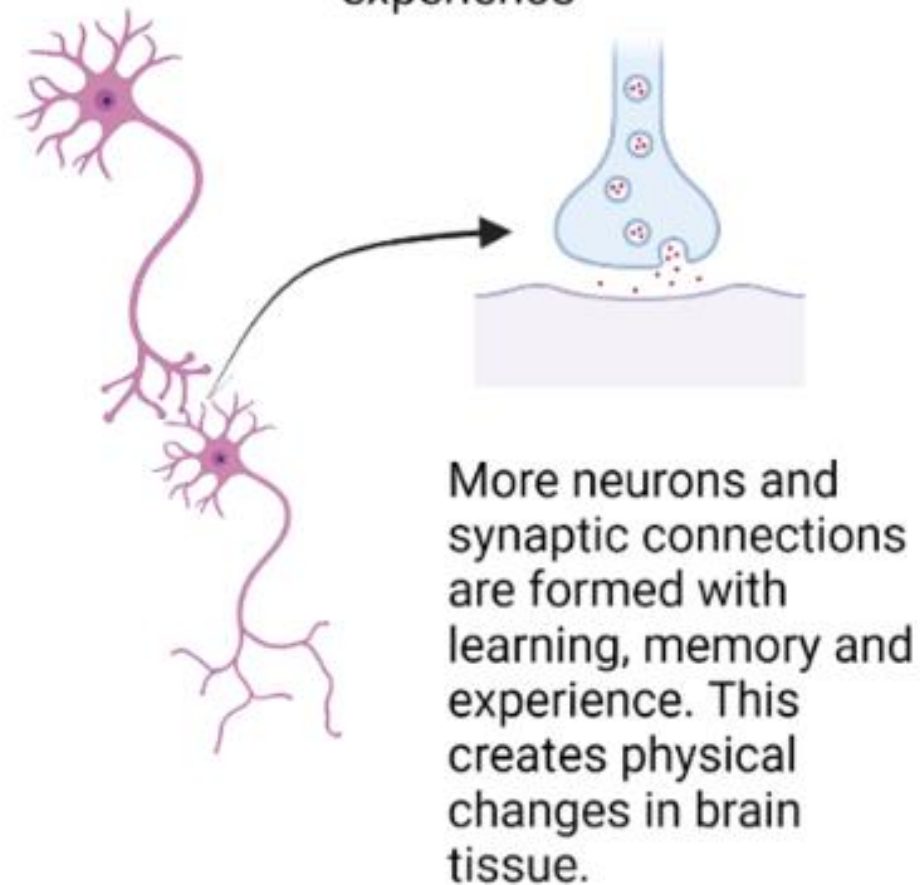
Pharmaceutical potential for patients with neuropsychiatric conditions



Communication pathways after (a) placebo and (b) psilocybin.

Structural Neuroplasticity

Ability of brain to physically change brain structure due to learning and experience

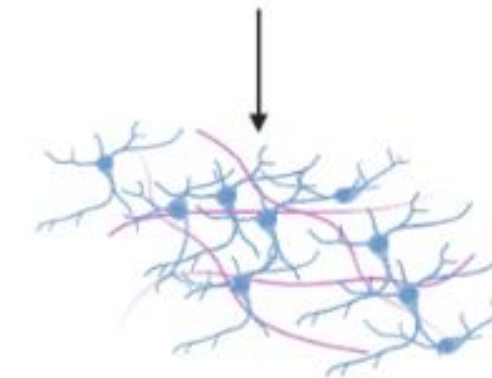


Functional Neuroplasticity

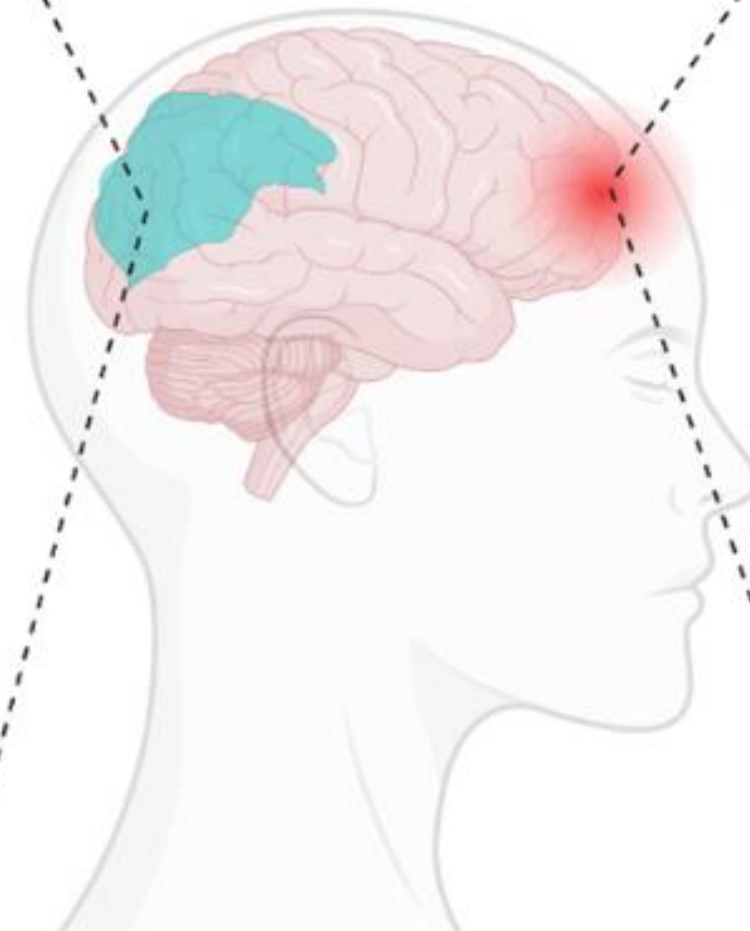
Synaptic remodeling following brain dysfunction or injury



Pre-existing neurons begin to die.



New neurons form synaptic connections over time to recover lost functions.



The immense clinical potential of Psilocybin treatment

Existing Clinical Programs include:

Treatment Resistant Depression [TRD]

Major Depressive Disorder [MDD]

Post-partum Depression

Post-Traumatic Stress Disorder [PTSD]

Obsessive Compulsive Disorder [OCD]

Depression in Bipolar-2 Disorder

Generalized Anxiety Disorder [GAD]

Body Dysmorphic Disorder [BDD]

Anorexia Nervosa

Binge Eating Disorder [BED]

Fibromyalgia Syndrome [FMS]

Irritable Bowel Syndrome [IBS]

Phantom Limb Pain

Migraine

Cluster Headache

Concussion Headache

Methamphetamine Use Disorder

Cocaine Use Disorder

Alcohol Use Disorder

Gambling Addiction

Smoking Cessation/Nicotine Addiction

Demoralization

Cancer-related mood & anxiety disorders

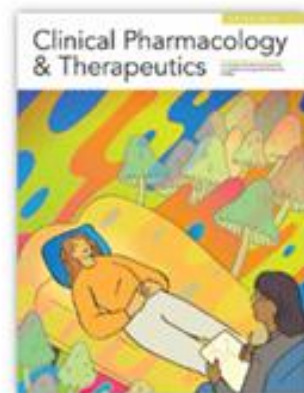
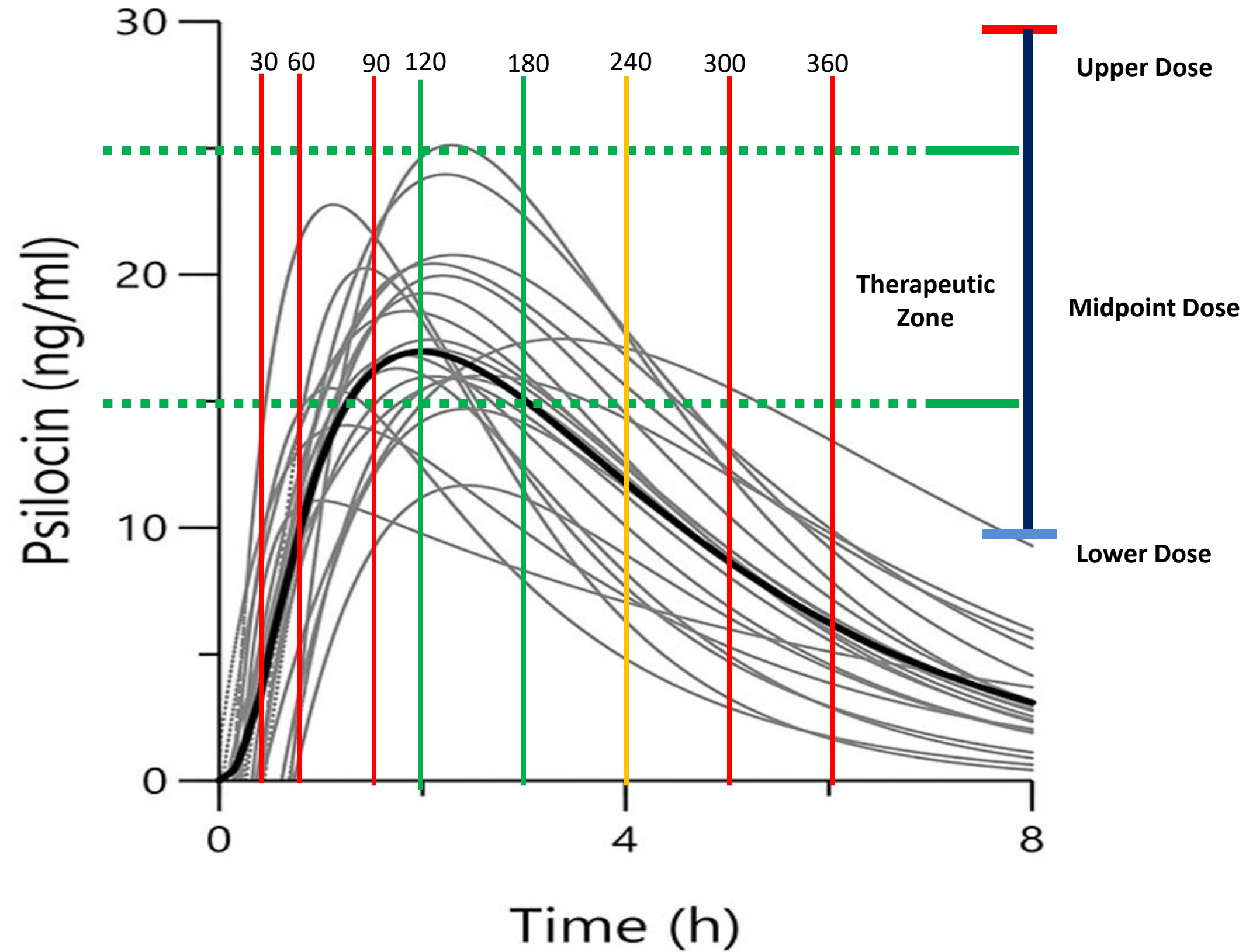
But how can the the clear clinical potential of Psilocybin be harnessed?

IV-infused Psilocin: A Precision Approach in Neuropsychiatry

	IV-infused Psilocin	Oral Psilocybin *
Short treatment duration of 1-2 hours	✓	✗ ~8-10 hours
Quick onset of psychedelic state (~15 minutes)	✓	✗ 1-2 hours
Precision targeting of drug blood levels in patients	✓	✗ highly variable
Quickly reversible in emergency	✓	✗
Strong IP positioning	✓	✗
Commercially scalable	✓	?

* Companies developing oral psilocybin include: Compass Pathways, USONA

Interpatient variability of oral psilocybin 25mg

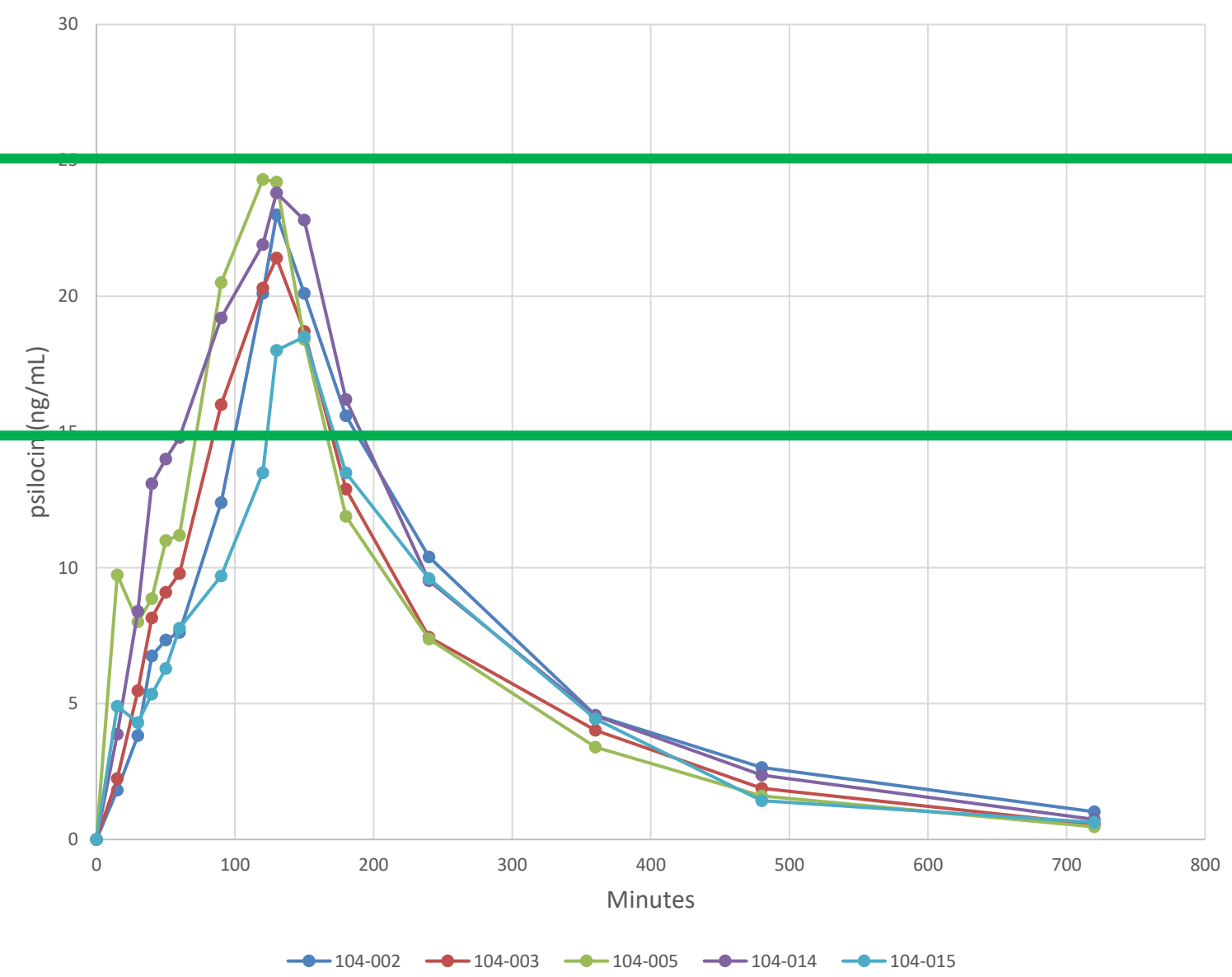
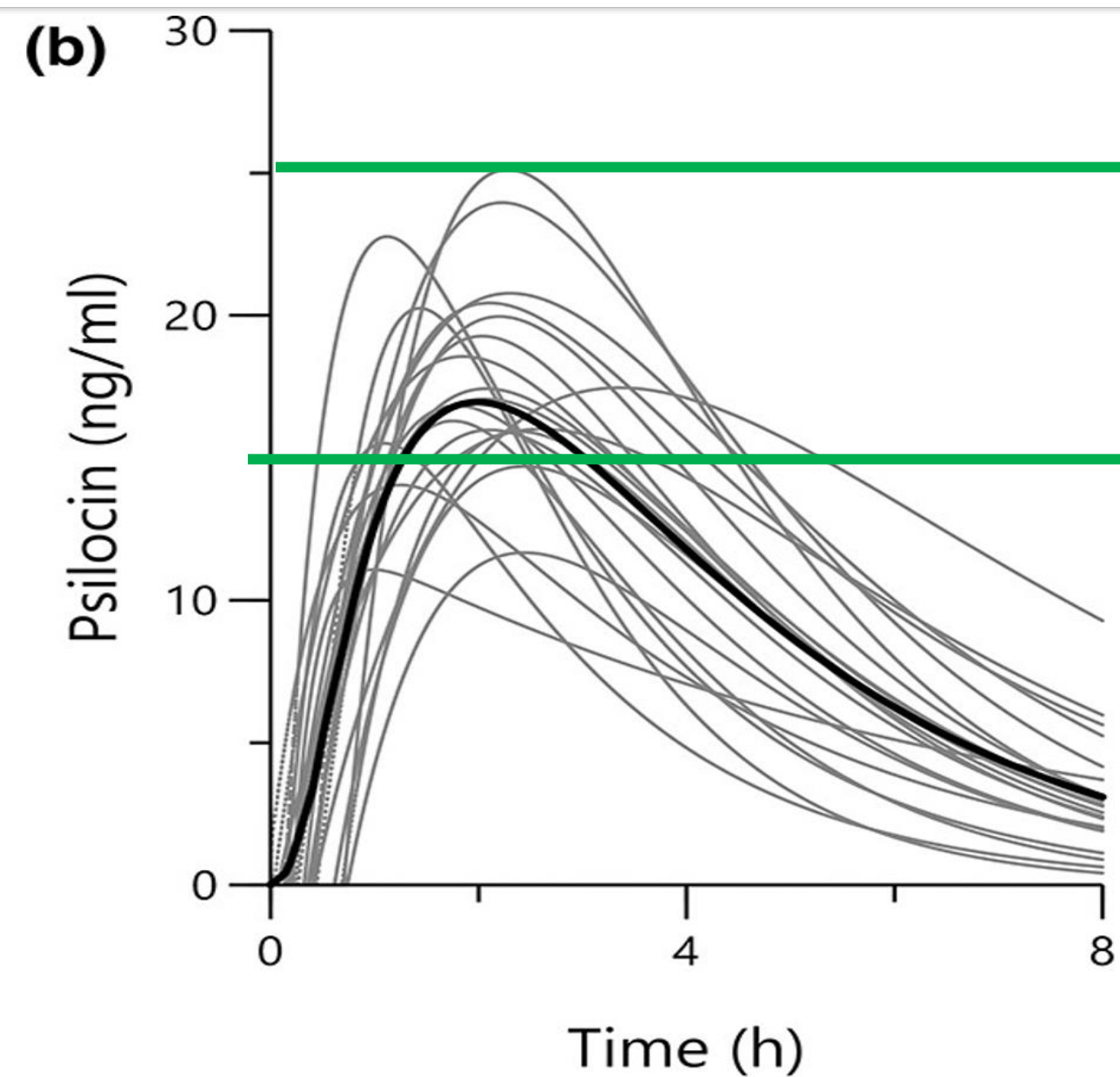


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Pages 747-751

Comparison of Psilocin Blood Level Variability: Oral Psilocybin vs TRP-8803 IV Infusion

Oral Psilocybin

TRP-8803



Clinical Trial Pipeline

Current
 Next 12-18 months**

TRP-8803	INDICATION	PHASE 1*	PHASE 2	
Proprietary IV-infused synthetic psilocin	BINGE EATING DISORDER			
	FIBROMYALGIA			
	IRRITABLE BOWEL SYNDROME			
TRP-8802	INDICATION	IND CLEARED	PHASE 2 a	PARTNER
Oral, synthetic psilocybin sourced from Usona Institute	BINGE EATING DISORDER			
	FIBROMYALGIA			
	IRRITABLE BOWEL SYNDROME			

*Healthy volunteer dose ranging study in 2024 will support IND submissions for Phase 2a studies in patients

**The timetable is indicative only and is subject to change.

Binge Eating Disorder (BED) : TRP 8802: Phase 2a clinical trial



Recurring episodes of eating large quantities of food and feeling unable to stop

25-50% of obese patients who seek weight-loss treatment suffer from problems with Binge Eating¹

No currently approved treatments developed specifically for Binge Eating Disorder

Patients suffering from BED have multiple co-morbidities²:

- 94% have lifetime Psychiatric disorders
- 70% Mood disorders
- 59% Depression
- 32% PTSD
- 23% of BED sufferers have attempted suicide

“We are very excited for the potential of TRYP’s treatment. The potential impact on patients’ lives is that it would be life changing for them.”

Jennifer Miller, MD, Professor, University of Florida, Principal Investigator

“These results from a single dose of psilocybin combined with therapy are clinically meaningful and highly promising. The magnitude of changes for most participants in binge eating, anxiety, and depression are dramatic..”

Jessie Dallery, Ph.D. Professor, University of Florida, Lead Psychologist

PRODUCT	NO. OF PATIENTS	COLLABORATOR	DESIGN	DATA READ OUT	NEXT STEPS
TRP-8802	6		Open label with psychotherapy	Data announced Q1 2023	Scientific paper publication

Positive interim data announced in January 2023, including mean reduction >80% for Binge Eating Score confirmed as viable target for future studies using TRP-8803

1. Bruce et al.; Journal of the ADA, Volume 96, Issue 1, Jan 1996, PP 58-61, Binge Eating Among the Overweight Population: A Serious and Prevalent Problem
 2. Keski-Rahkonen: Current Opinion in Psychiatry 34(6):p 525-531, November 2021. Epidemiology of Binge Eating Disorder: prevalence, course, comorbidity & risk factors

University of Florida Phase 2a BED Study Analysis:

Significant reduction in frequency & extent of binge eating (daily, 4 weeks)

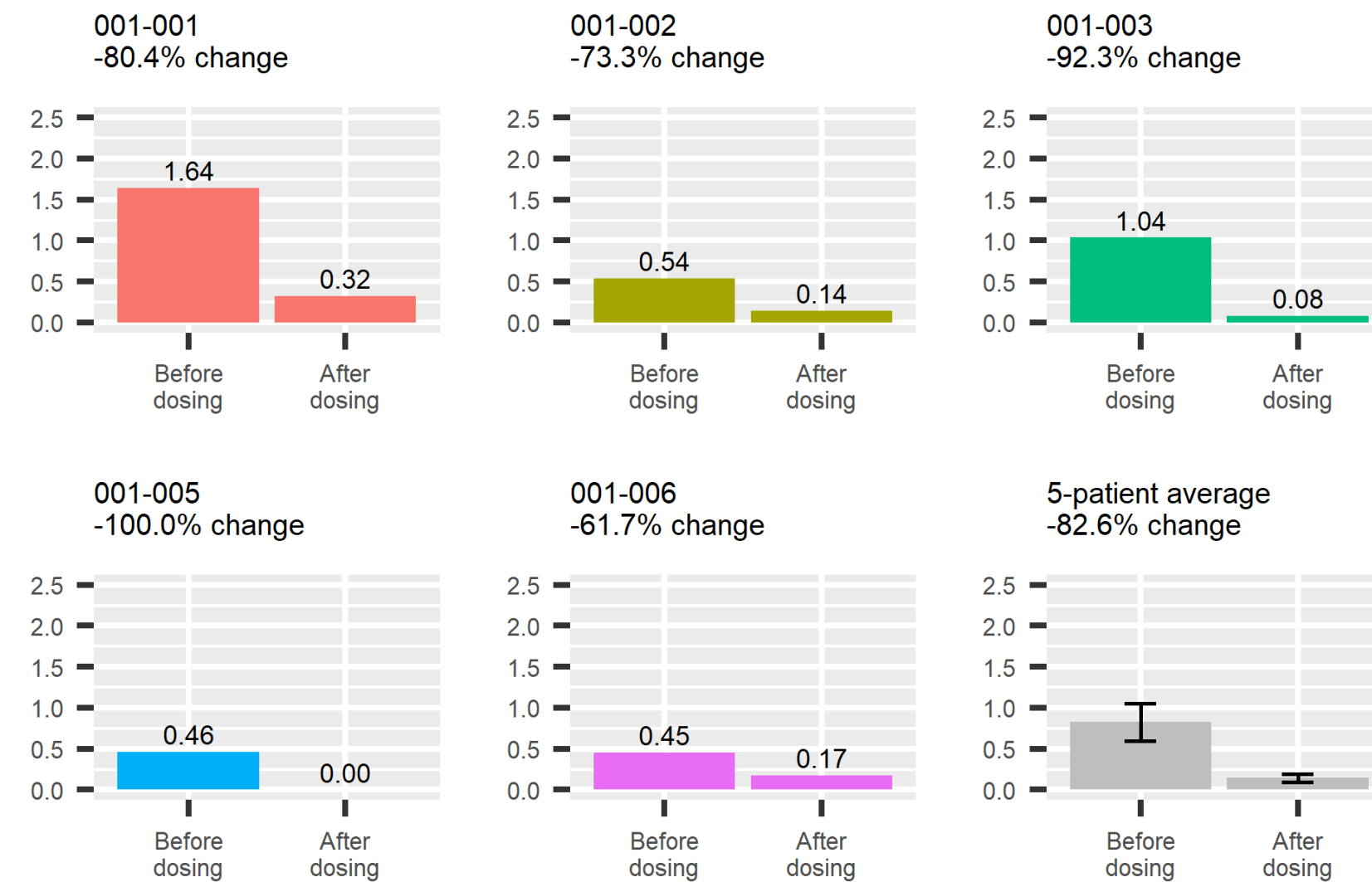
Question 1: Over the past 24 hours, how many times have you eaten what other people would regard as an unusually large amount of food (given the circumstances)?

Number of events per day (average)



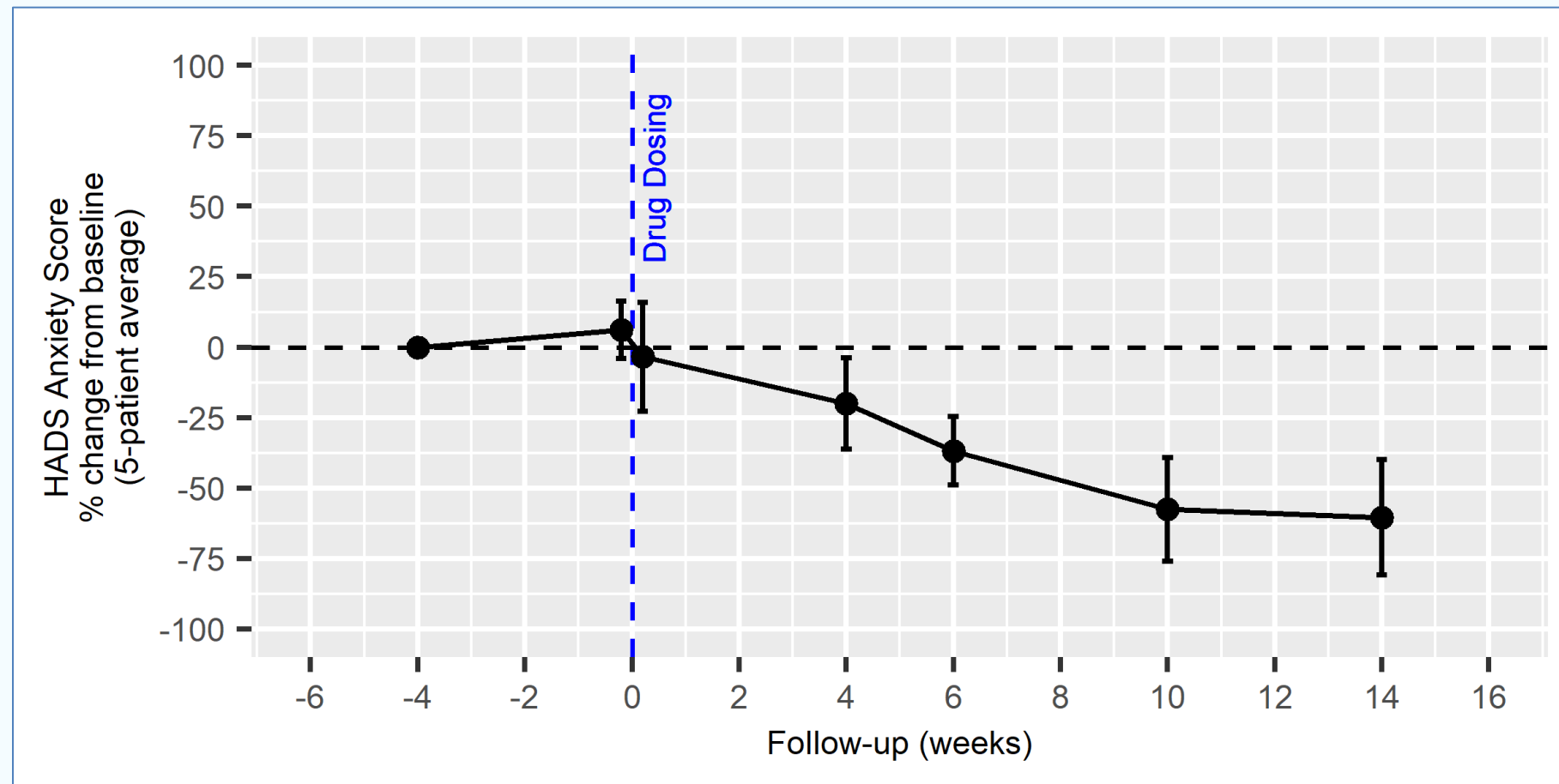
Question 2: On how many of these times did you have a sense of having lost control over your eating (at the time that you were eating)?

Number of events per day (average)

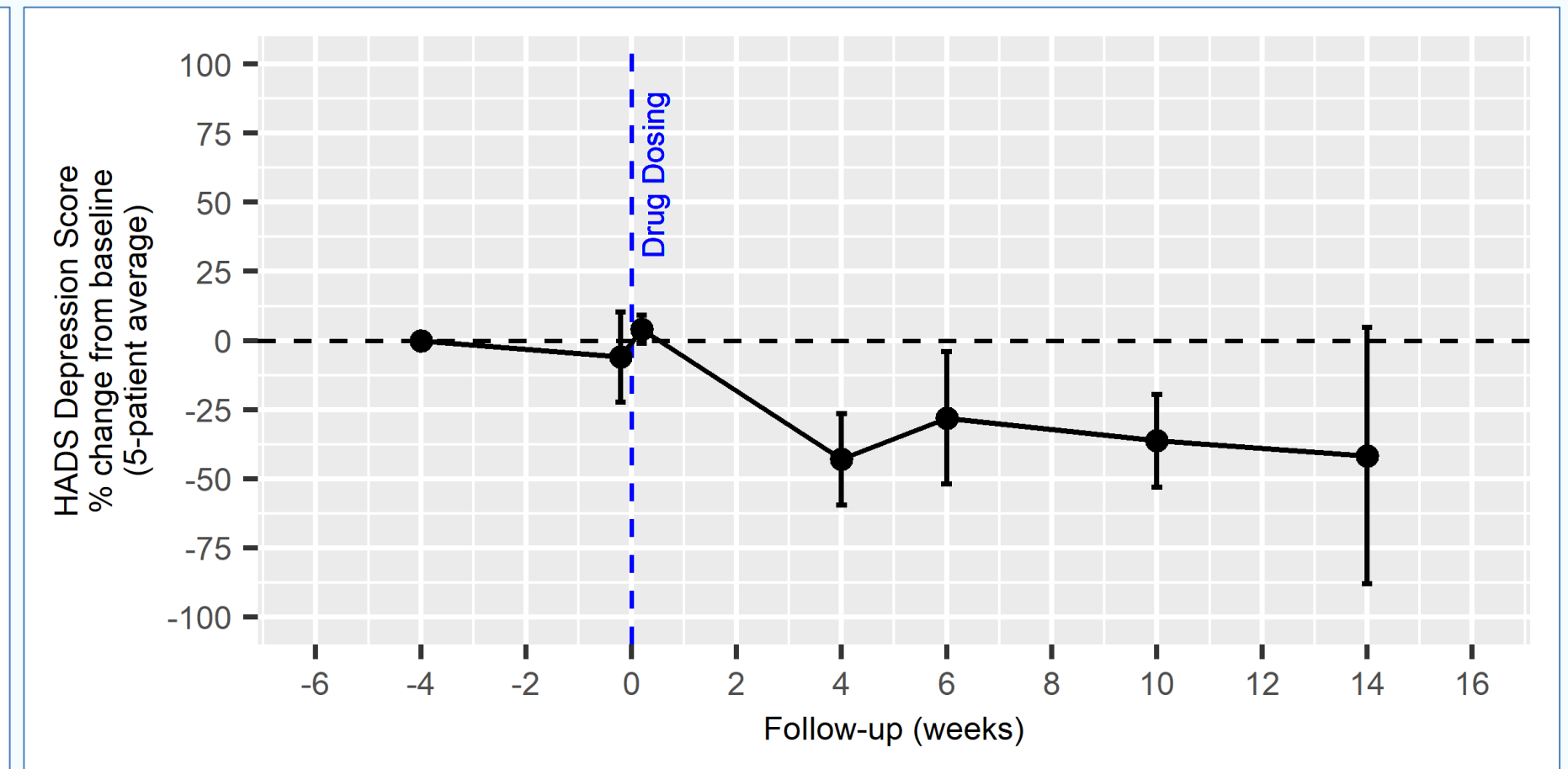


University of Florida Phase 2a Study Analysis:

Significant fall in Hospital Anxiety & Depression Scores (HADS)



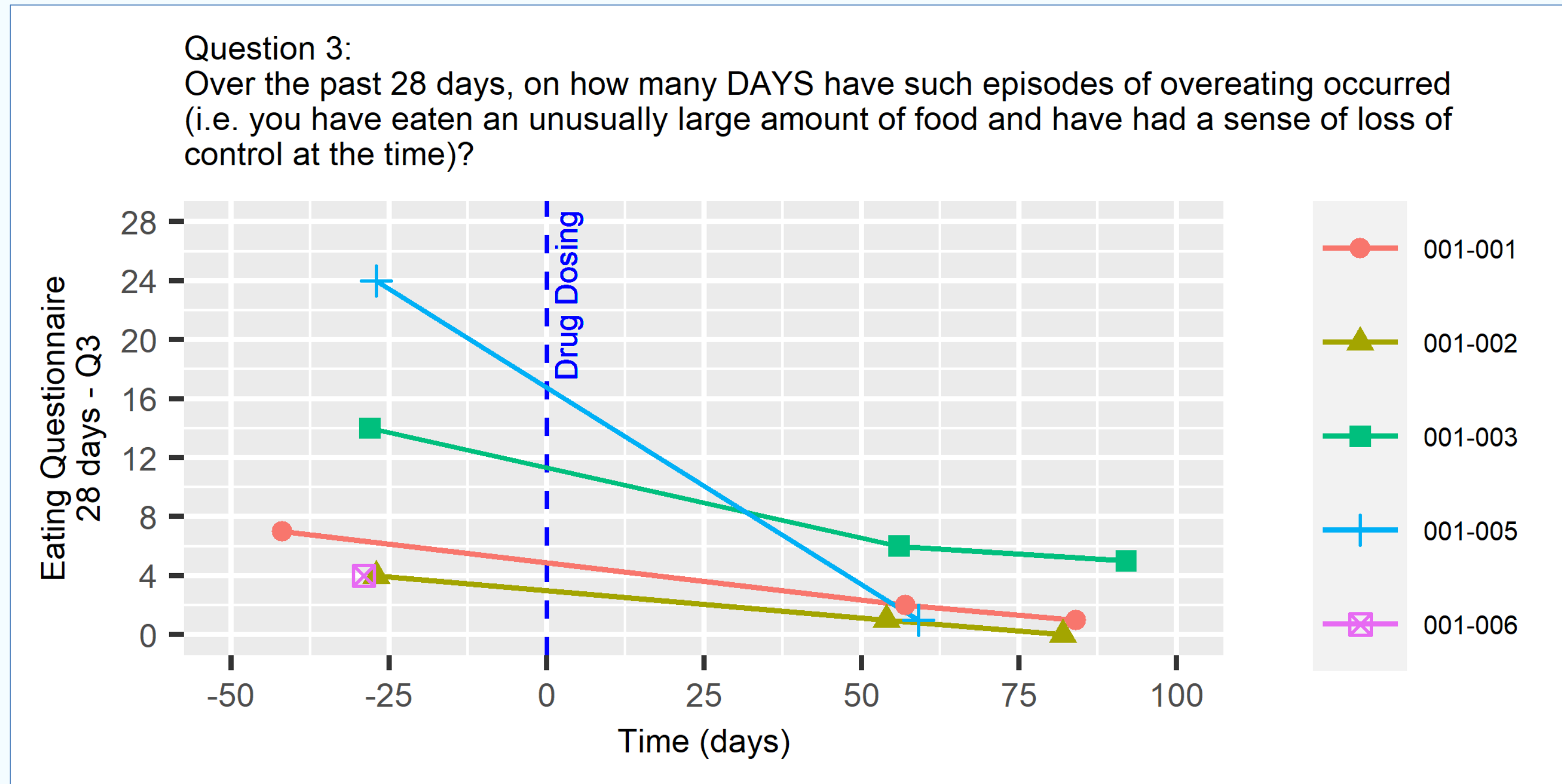
HADS Anxiety Score



HADS Depression Score

University of Florida Phase 2a Study Analysis:

Durable effect on binge eating episodes



Fibromyalgia: TRP 8802: Phase 2a clinical trial

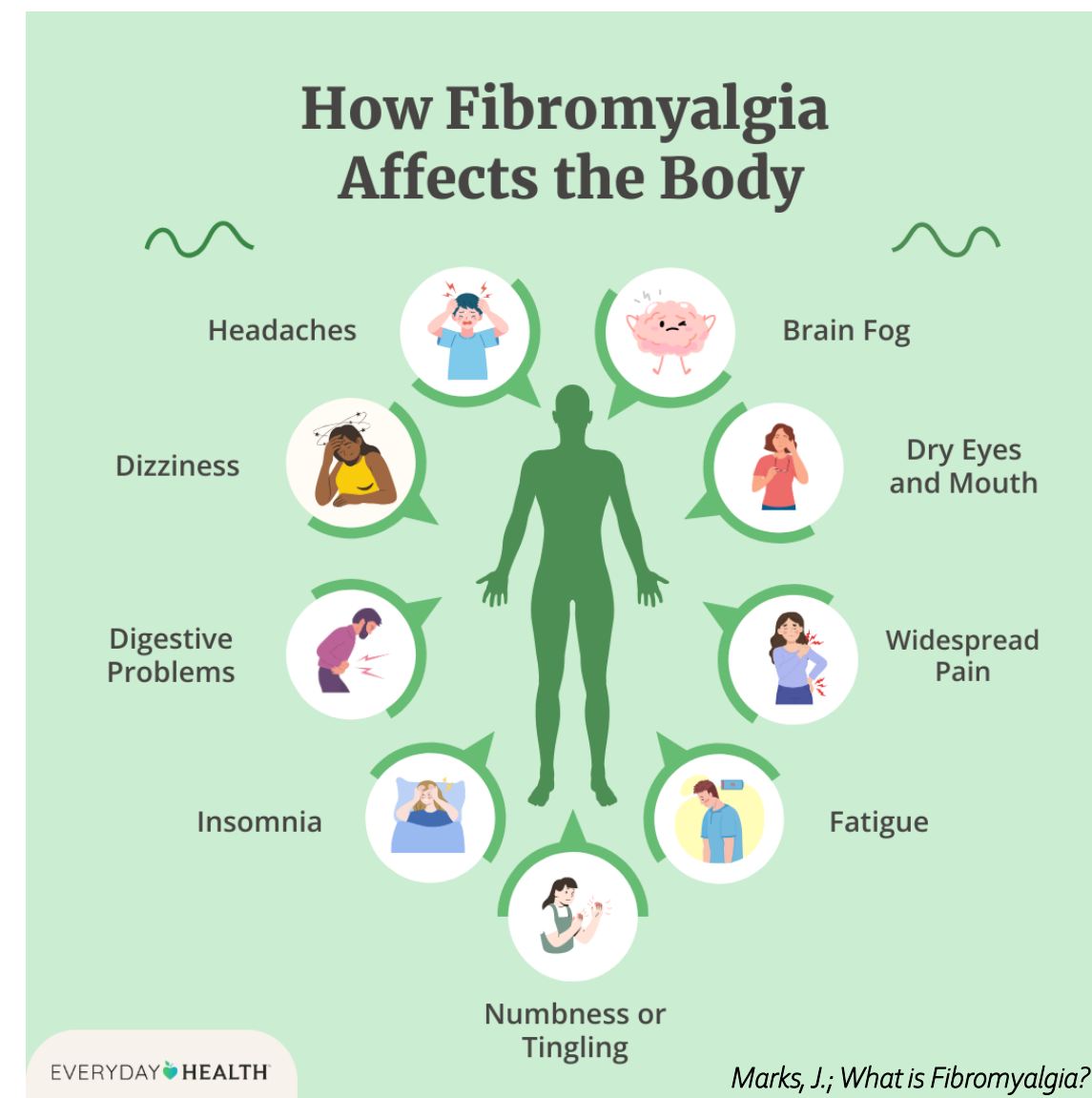
FMS characterized by widespread musculoskeletal pain, profound fatigue, sleep disturbances, and numerous other symptoms¹

Symptoms of fibromyalgia often begin after physical or emotional trauma, such as an illness, surgery, infection, life event or injury²

While fibromyalgia pain feels like it's coming from a specific area of your body, it's actually originating in your brain, specifically from the nervous system²

Many drugs have a limited effect on Fibromyalgia Pain¹

Co-morbidities include depression and health-related anxiety, sleep disturbances and increased suicide risk²



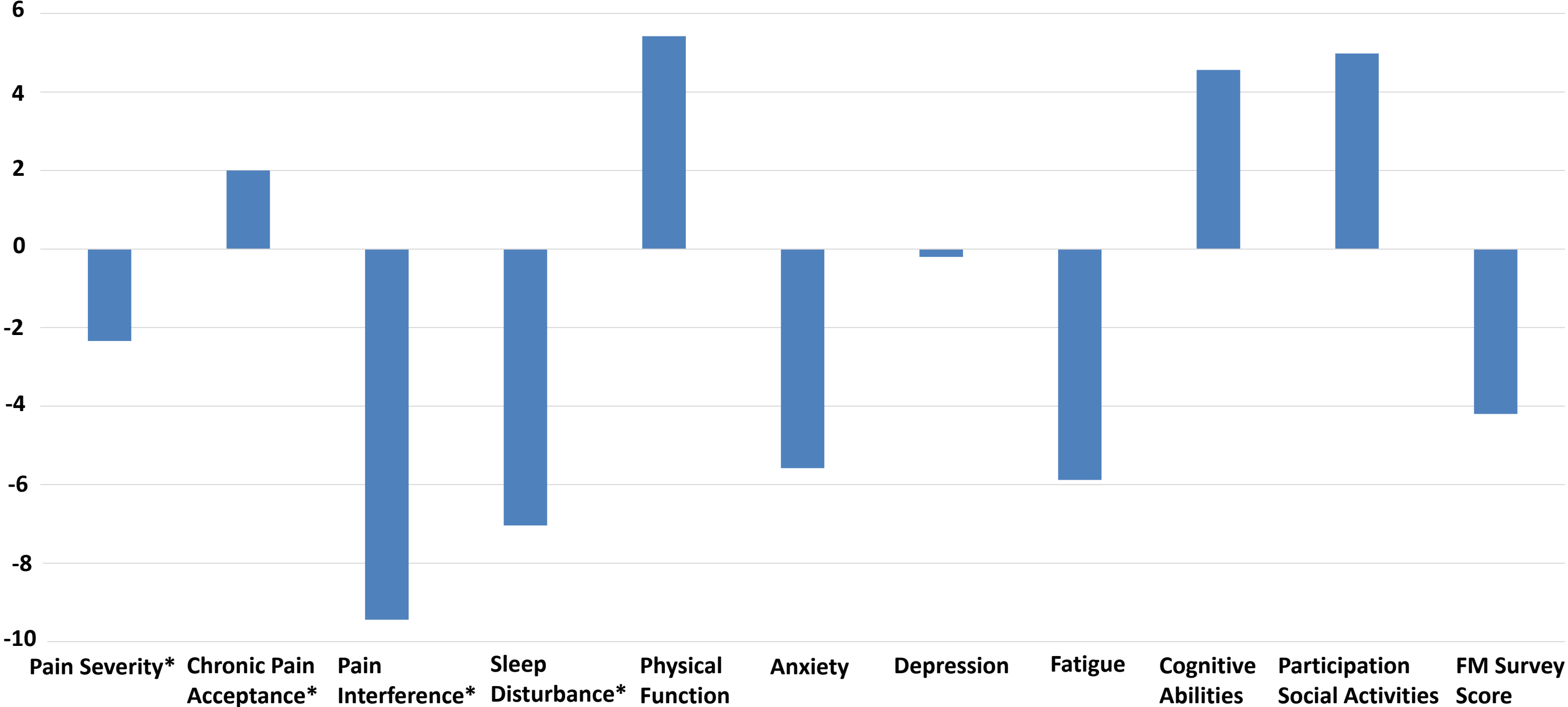
PRODUCT	NO. OF PATIENTS	COLLABORATOR	DESIGN	DATA READ OUT	NEXT STEPS
TRP-8802	Up to 10	UNIVERSITY OF MICHIGAN	Open label with psychotherapy	Initial Data Available	Full Clinical Study Data Release & 3 month follow up

First patient dosed in December 2023 with Data presented August 2024

1. Giorgi et.al.; Current Pain & Headache Reports; 23 July 2024; Pharmacological Treatment of Fibromyalgia Syndrome: A Practice-Based Review
2. Marks, J.; What is Fibromyalgia? Symptoms, Causes, Diagnosis, Treatment & Prevention; Everydayhealth.com/fibromyalgia/guide; Dec 15 2022

Change in Fibromyalgia symptoms following TRP-8802 administration:

(Baseline vs. end of treatment) for n=5 participants



For T-scores, changes of 2-6 points are considered a meaningful change. For pain severity, a 2-point difference is considered clinically significant. <https://www.healthmeasures.net/score-and-interpret/interpret-scores/promis/meaningful-change>

*Indicates secondary Outcome. CPAQ: Chronic Pain Acceptance Questionnaire. Pain Severity reported as change in aggregate pain score from the 7 days prior to the intervention to the end of the intervention. Sleep disturbance, pain interference, physical function, anxiety, depression, fatigue, participation in social activities, and cognitive abilities are all reported as T-scores per PROMIS scoring. Negative change scores indicate improvement for pain severity, pain interference, sleep disturbance, FM score, anxiety, depression, and fatigue. Positive change scores indicate improvement for CPAQ, physical function, participation in social activities, and cognitive abilities.

Irritable Bowel Syndrome (IBS) : TRP-8802 : Phase 2a clinical trial



Chronic abdominal pain + altered bowel habits

Affects 10-15% worldwide (~790M people)¹; leading cause of work absenteeism²

Associated with fibromyalgia, chronic fatigue, depression & anxiety

More common in those with early life adversity/trauma

Pathophysiology: visceral hypersensitivity

90% of serotonin synthesized in gut; enteric nervous system³

“There is tremendous potential for the treatment of debilitating IBS and other disorders of gut-brain interaction by utilizing the combined administration of psilocybin and psychotherapy. Our clinical study will examine how psilocybin-assisted psychotherapy may alter important brain networks involved in chronic pain and gastrointestinal-specific anxiety in IBS to bolster the neural flexibility in these patients and thereby reduce visceral hypersensitivity”

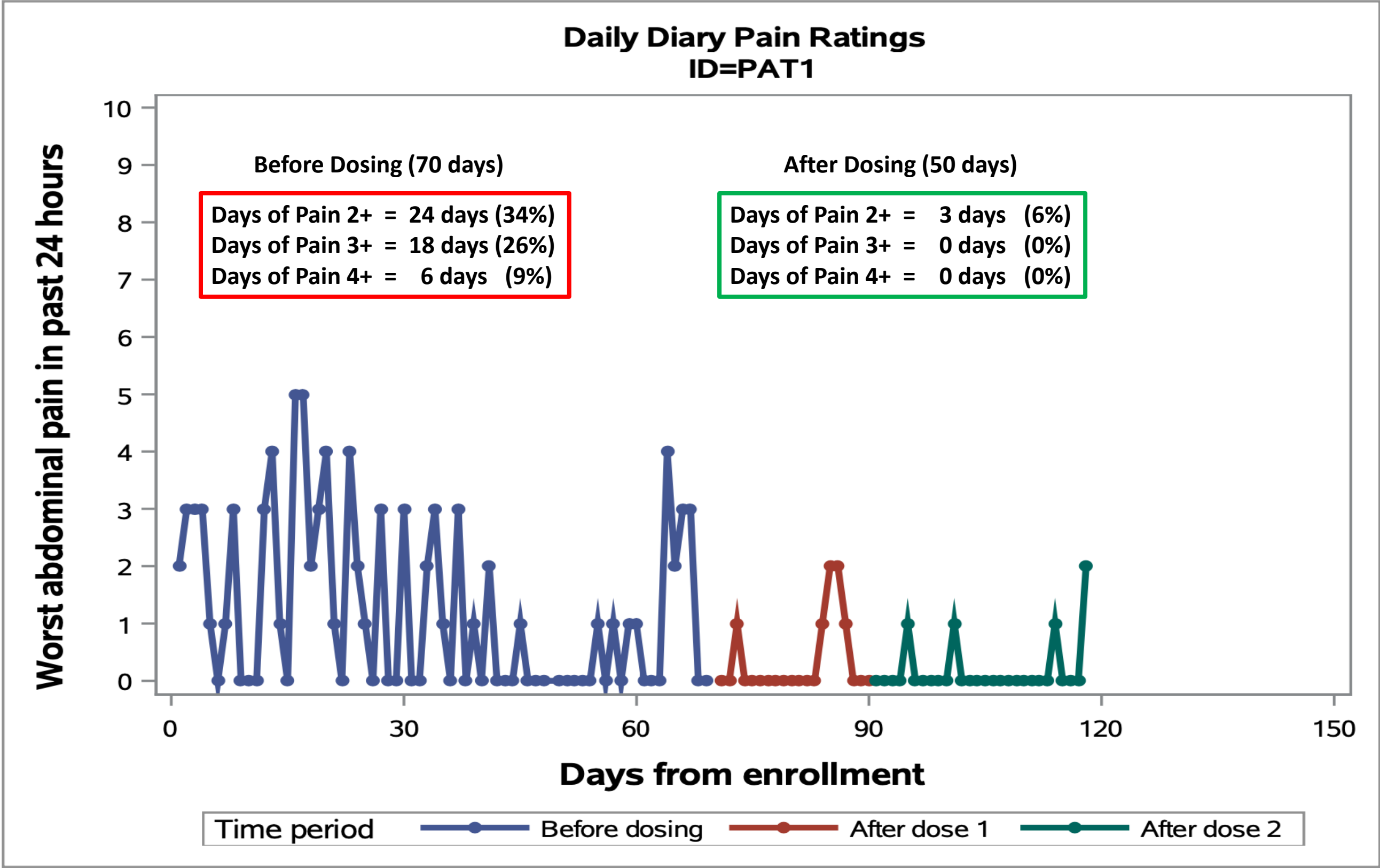
Erin Mauney, MD, Massachusetts General Hospital

PRODUCT	NO. OF PATIENTS	COLLABORATOR	DESIGN	DATA READ OUT	NEXT STEPS
TRP-8802	Up to 10	 MASSACHUSETTS GENERAL HOSPITAL	Open label with psychotherapy	H2 2024	Dose completion

Four patients dosed July – Sept 2024; Next steps Dosing Completion

- <https://gi.org/topics/irritable-bowel-syndrome>
- <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5010380/>
- <https://www.nature.com/articles/s41598-022-05756-0>

Patient 1 pain diary:



*Patient 1 diary-reported scoring of abdominal pain severity (out of 10) from Phase 2a Interim Analysis – MGH

TRP-8803: Healthy Human Volunteer Study (Australia):

A global first study now complete

IV-Infused Psilocin

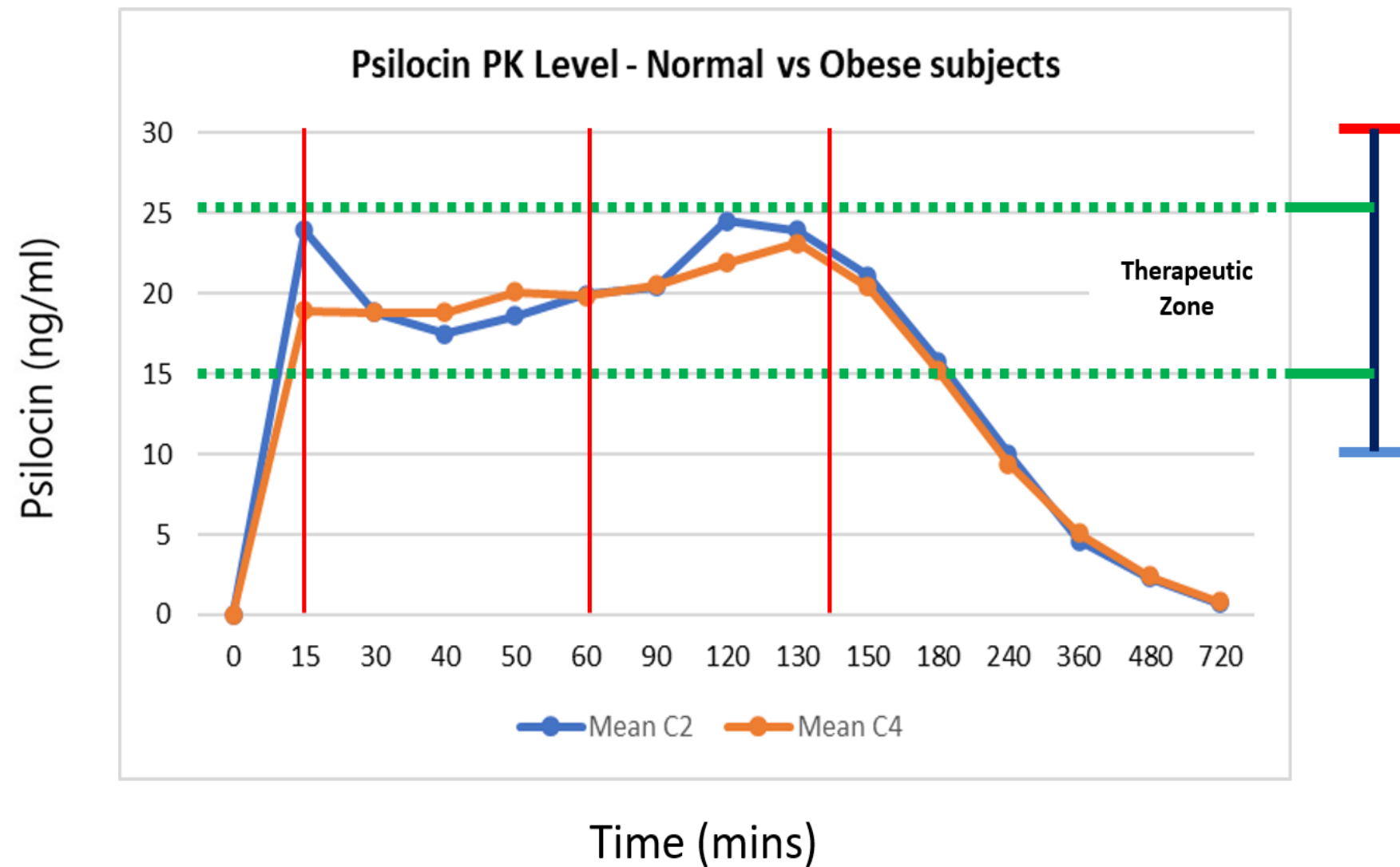
- Precision targeting of psilocin blood levels
- Fast onset time to psychedelic state (less than 20 mins)
- Short duration of treatment compared with oral psilocybin
- No weight-based dosing required
- Fully reversible in emergency
- IP protection
- Commercially scalable
- Enables clinical studies in multiple indications
- Safety Review Council finds TRP-8803 safe & well tolerated

HREC approved trial completed August 2024

PRODUCT	NO. OF PATIENTS	COLLABORATOR	DESIGN	DATA READ OUT	NEXT STEPS
TRP-8803	14 incl. 3 obese	iNGENu, C-Max	Open label with therapist support	H2 2024	Safety Review Council Report

Preliminary Results Phase I TRP-8803 Clinical Safety & Dose Ranging Study

TRP-8803 Achieves Consistent Blood Levels between Cohorts



- Established Safety of TRP-8803
- Confirmed Ability to achieve target blood levels of psilocin
- Confirmed reversibility of TRP-8803
- Achieved desired PK profile to reduce Side Effects while
- Established doses and infusion rates to be tested in Phase 2 Patient Studies

Robust Intellectual Property Portfolio

Patent applications and trade secrets based on novel methods for manufacturing, formulation, dosing, and specific disease indications

- Filed a provisional patent in March 2021 (US 63/161,070) covering TRP-8803 (IV-infused Psilocin); converted to PCT filing March 2022; published September 22, 2022
- Provisional patent application covering the use of psilocybin in the treatment of Binge Eating Disorder (BED) filed June 2022
- Provisional patent application for the treatment of fibromyalgia submitted September 2022
- Provisional patent application for salt & co-formers of TRP-8803 filed September 2022
- Provisional patent for IBS filed January 2, 2023

MORRISON
FOERSTER

Allens > Linklaters

Multiple Near-Term Milestones and Catalysts*

Catalyst	Timeframe	Status
Completion of Tryp Therapeutics Inc. acquisition	H1 2024	✓
\$6.5m capital raise	H1 2024	✓
Resumption of trading on ASX	H1 2024	✓
Appointments to strengthen Scientific Advisory Board	H1 2024	✓
Start of TRP-8803 Phase 1 trial (Australia)	H1 2024	✓
TRP-8802 Fibromyalgia Phase 2a patient enrolment (in collaboration with University of Michigan)	H1 2024	✓
TRP-8802 Irritable Bowel Syndrome (IBS) Phase 2a trial commencement (alongside Harvard University)	H2 2024	✓
Completion of TRP-8803 Phase 1 trial (Australia) and interim results	H2 2024	✓
TRP-8802 Fibromyalgia Phase 2a interim data	H1 2025	✓
TRP-8802 IBS Phase 2a interim data	<i>H2 2024</i> H1 2025	✓
TRP-8802 Fibromyalgia Phase 2a final data	H1 2025	
TRP-8803 Phase 2 trial <u>authorisations</u>	H1 2025	
TRP-8803 Phase 2 trial eating disorder trial commencement (Australia)	H1 2025	
TRP-8802 IBS Phase 2a final data	<i>H1 2025</i> H2 2025	
Commencement of TRP-8803 Phase 2 Binge Eating Disorder [BED] Trial (Australia)	H1 2025	
Commencement of TRP-8803 Phase 2 Fibromyalgia (chronic) pain Trial (Australia)	H2 2025	

*The timetable is indicative only and is subject to change (Calendar year is used)

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